

# THE IMPACT OF ENVIRONMENTAL TOBACCO SMOKE EXPOSURE ON CARDIORESPIRATORY FITNESS IN CHILDREN: A PILOT STUDY

MELISSA PARNELL, IVAN GEE, LAWRENCE FOWEATHER, GREG WHYTE,  
ZOE KNOWLES & JOHN DICKINSON

Public Health Institute & Physical Activity Exchange, Liverpool John Moores University, United Kingdom.

## ABSTRACT

Environmental tobacco smoke (ETS) in indoor air is a substantial risk factor for many health issues. Children are particularly susceptible to ETS with increased risk of asthma attacks, respiratory infections, and sudden infant death syndrome. The health effects of ETS are well researched in adults, but few studies examine the impact on children's cardiorespiratory fitness (CRF). CRF has been shown to be a useful biomarker for monitoring health effects which would normally be too subtle to identify at rest. In adults, ETS has been shown to reduce CRF, and children may be at greater risk due to high respiration rates and developing organs. This preliminary research tests the hypothesis that ETS has a detrimental impact on CRF in children. Twenty-five children (9-11 yrs) from one Merseyside primary school were recruited. ETS exposure was determined by parental surveys and coupled with children's exhaled carbon monoxide concentration. CRF was determined using a  $\text{VO}_2$  peak test, with lung function assessed using standard spirometry, and fractional exhaled nitric oxide (FeNO) provided an indication of lung inflammation. Initial results show children exposed to ETS had statically lower CRF scores ( $p = 0.048$ ) and were more likely to be classified as 'unfit' compared to children not exposed. A negative correlation was found between the number of cigarettes smoked at home and children's CRF ( $r = -0.526$ ,  $p = 0.008$ ), suggesting a possible dose-response relationship. Spirometry and FeNO values were not statistically different between groups. Results indicate that ETS exposure is likely to be detrimental to children's CRF. They highlight the need for further work, on a larger dataset that will allow more robust analysis with greater statistical power. To the author's knowledge, this study is the first of its kind to use laboratory-based fitness measurements to explore associations between ETS and CRF in children.

*Keywords: environmental tobacco smoke, cardiorespiratory fitness, children, spirometry, FeNO, tobacco, indoor air pollution.*

## 1 INTRODUCTION

Approximately 83% of tobacco smoke is in an invisible and gaseous form [1] which inadvertently exposes non-smokers to the effects of environmental tobacco smoke (ETS), often referred to as second-hand smoke (SHS), a common and significant indoor air pollutant. The effects of second-hand smoking are substantial and rapid, with effects predicted to be almost as large as chronic active smoking [2]. In adults, ETS can cause coronary heart disease, stroke and lung cancer, and exposed children can suffer from numerous health problems including severe asthma attacks, respiratory infections, ear infections and sudden infant death syndrome [3]. Like all carcinogens, there is no risk-free level of SHS exposure [4]. Worldwide, approximately 34% of non-smoking adults and 40% of children are exposed to second-hand smoke [5]. Öberg et al. calculated that in 2004, 603,000 deaths (approximately 1% of worldwide mortality) were attributed to ETS, and 10.9 million disability adjusted life years. Two main constituents of cigarette smoke, nicotine and carbon monoxide (amongst others) exhibit toxic effects on cardiovascular function both at rest and during exercise in adults [6]. The risk from ETS exposure is time dependant, and therefore limiting exposure early on is the most effective strategy for a reduction in tobacco smoke morbidity and mortality [7]. Children are particularly susceptible to the effects of ETS due

to their high respiratory rates and immature organs [8] and therefore extra effort should be taken to safeguard children from second-hand tobacco smoke.

Physical fitness is an integrated measure of most body functions involved in the performance of daily physical activity. Therefore, the functionality of all these systems is being checked when physical fitness is tested [9]. Cardiorespiratory fitness (CRF) is a health-related component of physical fitness defined as the ability of the circulatory, respiratory, and muscular systems to supply oxygen during sustained physical activity [10]. The maximal oxygen consumption ( $\text{VO}_2\text{max}$ ) attained during a graded maximal exercise to voluntary exhaustion has long been considered by the World Health Organisation as the single best indicator of CRF [11]. There is robust evidence that poor CRF is an independent risk factor for obesity, diabetes, and heart disease related mortality, and that a moderate to high level of CRF reduces the risk of all-cause cardiovascular disease (CVD) mortality [12], [13].

Subtle changes in the function of the cardiorespiratory system can be recognised through exercise-based measures. Therefore, exercise related health markers, such as  $\text{VO}_2\text{peak}$ , may provide novel methods for measuring health in relation to ETS exposure in children. To date, no previous research has sought to examine the impacts of ETS exposure on children's laboratory measured  $\text{VO}_2\text{peak}$  and therefore the present study will provide new insight to current body of knowledge. Using a small initial sample population, this exploratory study aims to examine the hypothesis that exposure to the indoor air pollutant, ETS, is detrimental to children's cardiorespiratory fitness.

## 2 METHOD

### 2.1 Recruitment

Prior to recruitment, ethical approval was obtained from Liverpool John Moores University Ethics Committee. Children from one Liverpool (United Kingdom) primary school, ages 9-11 years ( $n = 25$ ) agreed to take part in the study. Inclusion criteria were children aged 9-11 years old and attend a Merseyside primary school. Exclusion criteria were any medical conditions that limit a child's ability to walk or run on a treadmill, including heart conditions where vigorous exercise would put the child at risk.

### 2.2 Quantitative Surveys

Questionnaires to determine self-reported household smoking status, adapted from the Global Adult Tobacco Survey [14] were completed by a consenting parent or guardian. Questions determined the number of tobacco smokers in the home, as well as which rooms smoking occurred, and how many cigarettes were smoked each day. Participant (adult and child) demographic information was also obtained via the parental questionnaire, with postcodes used to determine English indices of multiple deprivation score (EIMD) [15]. Children completed a Physical Activity Questionnaire for Children (PAQ-C) to determine self-reported levels of physical activity.

### 2.3 Anthropometrics

Body mass to the nearest 0.1 kg (Seca, Birmingham, UK), stature and sitting stature to the nearest 0.1 cm (Seca, Birmingham, UK) were assessed using standard techniques [16]. Body mass index (BMI) was calculated from height and weight ( $\text{kg}/\text{m}^2$ ), with age and sex specific BMI cut-points used to classify child weight status [17].

## 2.4 Exhaled Gases

Exhaled carbon monoxide (eCO) was measured using a breath Smokerlyzer PiCO device (Bedfont, UK) to determine recent ETS exposure. Pilot work indicated that eCO quickly declined several hours post-ETS exposure. Therefore, eCO measurements were taken on the morning (08:30) of laboratory visits to better reflect ETS exposure from the home. Fractional exhaled nitric oxide (FeNO), an indication of airway inflammation, was measured at rest (twice) using a NIOX® VERO device (Circassia, UK) in the laboratory, prior to spirometry.

## 2.5 Spirometry

Spirometry measures; forced vital capacity (FVC), forced expiratory volume (FEV<sub>1</sub>), peak expiratory flow (PEF), forced expiratory ratio (FER) were taken at rest using a digital micro-spirometer (Micro-plus spirometer, CareFusion, UK). Spirometry measures were repeated at least three times with the highest value recorded.

## 2.6 Cardiorespiratory Fitness

Cardiorespiratory fitness (CRF), peak oxygen uptake (VO<sub>2peak</sub>), was assessed using an individually calibrated, continuous incremental treadmill test to volitional exhaustion using breath by breath analysis (Jaeger Oxycon Pro, Viasys Health Care, UK). This method is based on an established protocol described previously by Boddy et al [18]. VO<sub>2peak</sub>, as opposed to VO<sub>2max</sub> (plateau of oxygen uptake), was used as children often fail to reach plateau [19]. However, experiments have shown that 'true' VO<sub>2max</sub> values can be achieved in children without the need for plateau if test endpoints are met [20].

## 2.7 Analysis

Data was analysed using a combination of Microsoft Office Excel 2016 and IBM Statistical Package for the Social Sciences (SPSS) v24. Incomplete datasets and participants which did not reach peak VO<sub>2</sub> [18] were excluded from the analysis of CRF.

# 3. RESULTS

## 3.1 Participant characteristics

Of the 25 children who returned written parental informed consent and participant assent, only one participant did not complete the VO<sub>2peak</sub> fitness test. Participant response rate for the one participating school was 40.9%. It should be noted that the recruited number of participants from smoking households was very low (n=2 females, n=5 males) and so thorough analysis of these individual groups is limited. Both the school and participant's homes are urban within the Liverpool City air quality management area and are included in the 20% most deprived neighbourhoods in England. White British children accounted for 88% of the sample, with 8% Polish, and 4% Portuguese ethnicity.

Males were found to have significantly higher VO<sub>2peak</sub> scores than females ( $p = 0.042$ ). Due to the finding, the dataset was split by sex for certain aspects of the analysis. To allow for male and female datasets to be analysed together, children were classified as aerobically fit or unfit according to established thresholds for male and female children; 46.6 and 41.9 ml/min/kg for males and females respectively [21].

### 3.2 Breath eCO as a measure of environmental tobacco smoke exposure

A moderate, statistically significant Pearson correlation ( $r = 0.504$ ,  $p = 0.010$ ) was found between eCO concentration and parental self-reported number of cigarettes smoked per day. Children with self reported ETS exposure had a higher mean eCO of 1.57 ppm compared to those no exposed, 1.00 ppm, although this was not statistically significant ( $p = 0.219$ ).

### 3.3 Spirometry and FeNO

Table 1: Mean spirometry values expressed as percentage of normative values for sex, height and ethnicity. FVC – Forced vital capacity, FEV<sub>1</sub> – Forced expiratory volume in 1 second, PEF - Peak expiratory flow, FER – Fractional exchange ratio, FeNO – Fraction of exhaled nitric oxide.

Measure (mean)	Non-smoking household (n=18)	Smoking household (n=7)	T-test $p$ values
%FVC	97.7 (18.1)	92.7 (24.6)	0.580
%FEV <sub>1</sub>	86.0 (20.8)	88.9 (19.2)	0.754
%PEF	88.2 (31.3)	82.6 (22.1)	0.743
%FER	92.2 (10.3)	98.4 (9.62)	0.178
FeNO (ppb)	22.0 (19.6)	40.1 (47.0)	0.178

No statistically significant differences were found for spirometry measures between exposure groups (Table 1). Mean fractional nitric oxide concentration in exhaled breath (FeNO) in the smoking group was almost double that of the non-smoking group (22 ppb and 40.1 ppb respectively), but this was not found to be statistically significant ( $p = 0.178$ ). The range for FeNO measurements was very high (140) with a minimum value of 5 ppb and a maximum of 145 ppb.

### 3.4 VO<sub>2</sub>peak between exposure groups

Fitness (VO<sub>2</sub>peak) was found to be significantly lower in the group exposed to ETS with a mean VO<sub>2</sub>peak of 50.64 and 44.64 (ml/min/kg) respectively for children exposed and not exposed to ETS ( $p = 0.048$ ). However, when the same analysis was performed separately for males and females, only the difference between the male exposure groups was significant (Table 2).

Table 2: Mean VO<sub>2</sub>peak scores by sex for children exposed and not-exposed to ETS in the home.

	Mean VO <sub>2</sub> peak $\pm$ SD		Statistical significance
	Exposed (n=7)	Not exposed (n=17)	
<b>Male</b>	46.28 (6.95)	55.47 (3.96)	$p = 0.015$
<b>Female</b>	40.55 (8.27)	47.25 (4.69)	$p = 0.125$

A moderate ( $r = -0.470$ ) yet significant ( $p = 0.021$ ) negative correlation (Pearson) was found between eCO and VO<sub>2</sub>peak when both sexes were analysed together (Fig. 1). The same significance was not found when the data was split by sex. A negative correlation was also found between parental self-reported number of cigarettes smoked per day and VO<sub>2</sub>peak,  $r = -0.526$ ,  $p = 0.008$ . Both sexes showed a reduction in fitness as the number of cigarettes smoked at home per day increased, with Pearson  $r = -0.635$ ,  $p = 0.026$  for males, and  $r = -0.630$ ,  $p = 0.028$  for females. The correlation (Pearson) no longer held statistical significance when the extreme value at 65 'cigarettes smoked per day' was removed,  $r = -0.314$ ,  $p = 0.145$ . However, when split by sex, the trend was still apparent for males,  $r = -0.635$ ,  $p = 0.026$ .

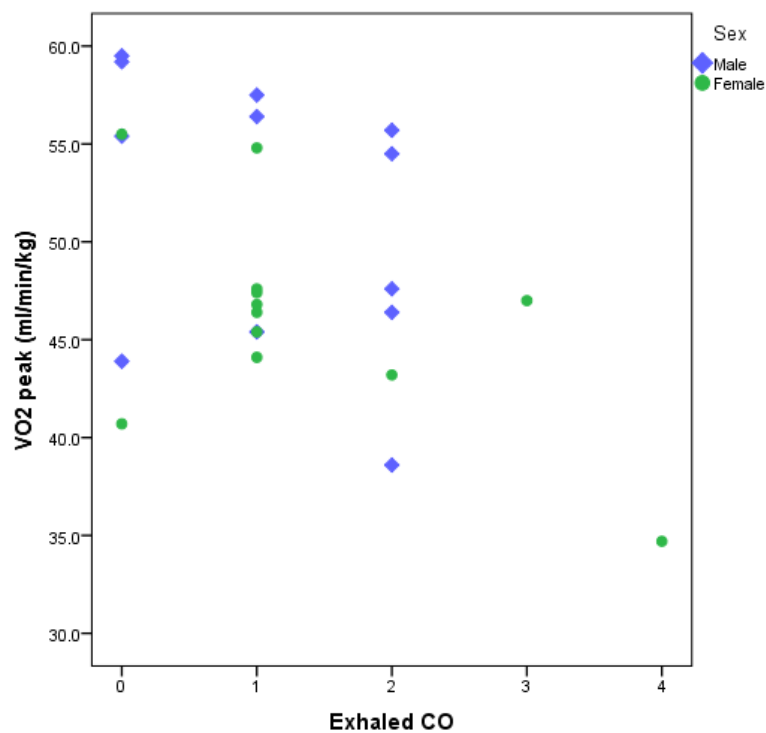


Figure 1: VO<sub>2</sub>peak (ml/min/kg) and eCO (ppm) for males and females.

More children were classed as fit from non-smoking households (94%) compared to smoking households (29%) where the majority were classed as aerobically unfit (Chi-square test,  $p = 0.001$ ), as shown in Table 3. Fitness thresholds were 46.6 and 41.9 ml/min/kg for males and females respectively.

Table 3: Percentage of children classed as fit from smoking (n=7) and non-smoking households (n=17).

	Fit (%)	Unfit (%)
<b>Smoking household (n=7)</b>	29	71
<b>Non-smoking household (n=17)</b>	94	6

### 3.5 Body Mass Index

A significant negative correlation (Pearson) was found between BMI and  $VO_2$ peak for both males ( $r = -0.876$ ,  $p < 0.001$ ) and females ( $r = -0.800$ ,  $p = 0.002$ ). The mean BMI was higher in the ETS exposed group, but this was only statistically significant for males (Table 4).

Table 4: Mean BMI scores for males and females from smoking and non-smoking households.

	Mean BMI $\pm$ SD		Statistical significance from t-test
	<b>Exposed</b>	<b>Not exposed</b>	
<b>Male</b>	20.05 (3.39)	16.78 (1.25)	$p = 0.044$
<b>Female</b>	20.17 (4.51)	19.00 (3.60)	$p = 0.786$

### 3.6 Deprivation

Deprivation score, EIMD, was negatively correlated with  $VO_2$ peak fitness score for males only ( $r = -0.683$ ,  $P = 0.020$ ). EIMD scores were found to be significantly higher for smoking families, 57.4, than non-smoking families, 41.2 ( $p = 0.023$ ).

### 3.7 Physical Activity

The mean self-reported physical activity (PA) scores were 3.59 (SD 0.65) for females and 3.46 (SD 0.70) for males (1 indicating low PA and 5 high PA). PA scores showed no correlation with  $VO_2$ peak scores ( $r = -0.001$ ,  $p = 0.997$  for males, and  $r = 0.087$ ,  $p = 0.780$  for females).

## 4 DISCUSSION

This initial study examined the association between exposure to ETS in the home, and children's CRF from a small sample of children from one Liverpool primary school. A significant difference in CRF was found between the children from smoking and non-smoking homes with 71% of children from smoking homes being classed as aerobically unfit,

compared to only 6% from non-smoking households. The mean CRF, as measured by  $\text{VO}_{2\text{peak}}$ , was 16% lower in male children from smoking households, and 6.7% lower in females from smoking households, compared to non-smoking households. However, 51% of children from smoking households were classed as being either overweight or obese, compared to 22% of children from non-smoking homes, which could be an important confounder.

Self-reported PA score was not found to be correlated  $\text{VO}_{2\text{peak}}$ , suggesting fitness is influenced by other factors. However, self-reporting surveys are notoriously unreliable, and further confounded by the fact the participants are children and may exaggerate their PA levels due to enthusiasm or desirability bias.

Exhaled CO was positively correlated with the number of cigarettes parents reported smoking at home each day. Both eCO and cigarettes smoked per day were negatively correlated with CRF in males and females. The results of the present study suggest that tobacco smoking, and the resulting CO inhalation, may be having a detrimental effect on children's CRF, as found previously in adults [22]. However, due to the nature of this study and the small sample size, causation cannot be assumed. ETS is not the only source of CO. The road transport microenvironment is a significant contributor to an individual's CO exposure, with travel by vehicle causing the highest level of CO exposure, compared to walking and cycling [23]. With this in mind, particle monitors (Dylos 1700) will be set up in homes and schools of smoking and non-smoking participating families to determine  $\text{PM}_{2.5}$  concentrations in the home. This information will be used to compare the air quality in smoking and non-smoking homes and to highlight any other significant PM exposures (e.g. traffic congestion near the home or school).

In the present study, FeNO values were highly variable with a range of 140, the lowest value at 5 ppb, and the highest at 145 ppb. Eosinophilic inflammation is likely at 35 ppb or higher [24]. Repeat measures on individuals only varied by 1-2 ppb, therefore the range is due to individual differences. No statistically significant associations were found between the exposure groups and FeNO, although the tobacco smoking group did have double the percentage of children classified as having a high concentration ( $>35$  ppb) of FeNO compared to the non-smoking group. There are a number of conditions that elevate levels of FeNO, including asthma and allergic rhinitis [25]. A larger dataset is needed to better determine whether elevated concentrations of FeNO are associated with ETS; this will be explored in future work.

Spirometry measures were expressed as a percentage of the normative values for age, height, sex, and ethnicity. Mean %FVC and mean %PEF were respectively 5.0% and 5.6% lower in the smoking group compared to the non-smoking group but this was not statistically significant. Parental smoking has been shown to reduce lung function in children, as measured through spirometry. For example, a small but significant decrease in  $\text{FEV}_1$  due to parental smoking and in utero exposure has been associated with reduced PEF and FVC [26], [27]. A major limitation with the use of spirometry with children is compliance; children often find it difficult to perform the test correctly and therefore recorded values may not reflect true ability. As described earlier, some subtle changes in system functions can only be picked up through exercise related measures. All spirometry measures in the present study were completed with the participant at rest, and further work will aim to include mid-flow spirometry measures during exercise which may be a more sensitive measure of lung function and health.

The present study shows that unfit males on average have higher deprivation (EIMD) scores. Children from more deprived households may have less opportunity for organised physical activity, which may explain the reduced  $\text{VO}_{2\text{peak}}$ , but it may be the case that

smoking habits, which are also associated with deprivation, are partly responsible for the difference. As EIMD scores are calculated using postcodes, and fitness was correlated to EIMD, local air quality may be also influencing children's fitness. Liverpool City is an Air Quality Monitoring Area (AQMA) with regular exceedances of national standards for nitrogen dioxide. However, the nearest automatic urban monitoring network (AURN) station is Speke, Liverpool, approximately 10 km from the participating school. Whilst this data could be utilised, it would not very well reflect the children's exposure from their microenvironment, especially not in the home. As stated above, further work will use PM<sub>2.5</sub> monitors to assess air quality both in the home and at the school, which will shed light on the contribution from other sources of air pollution, as well as indoor ETS.

The number of females from smoking households was extremely low in this sample (n=2), and therefore statistical analysis of this group is limited. Participants from smoking households accounted for 28% of the sample, with the majority reporting to smoke 20 or less cigarettes per day, and one participating family smoking at 65 per day. It would be useful to recruit children from households who smoke more than 20 cigarettes per day, which might elude to a linear or otherwise trend between child CRF and household cigarettes smoked per day. Whilst the ratio of males to females was almost 50%, white British children accounted for 88% of the participants in the current study and further work will include participants from a variety of ethnic backgrounds.

ETS poses a significant threat to an individual's health, with children at a greater risk due their developing organs. CRF is a good indicator of general health as it encompasses all of the body's major systems and organs [9]. An overweight and unfit child is likely to become an overweight and unfit adult [28], which is of great concern for the future of public health. The current study has provided an insight into ETS exposure and the potential detrimental impact on children's CRF. This study has demonstrated a significant difference between the two exposure groups with a small sample size. It will be of great interest and importance to see if these differences stand true with a larger and more diverse sample.

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